# MMR

## MORBIDITY AND MORTALITY WEEKLY REPORT

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# Use of Death Certificates for Surveillance of Work-Related Illnesses — New Hampshire

To conduct surveillance for occupationally related health events, some state health departments have made innovative use of the limited data sources available to them locally (e.g., death certificates, workers' compensation claims, hospital discharge files) (1). The New Hampshire Division of Public Health Services (NHDPHS) of the State Department of Health and Human Services recently completed a study of death certificates in the state for 1963-1983 to epidemiologically characterize deaths due to mesothelioma and the pneumoconioses (silicosis, asbestosis, and anthracosis).

Death certificates were provided by New Hampshire's Bureau of Vital Health Statistics for 1963-1983 and were reviewed to obtain information on all deaths for which mesothelioma or a pneumoconiosis was recorded as the underlying or contributing cause of death. Information was abstracted from these certificates on sex, occupation, age at death, year of death, and usual place of residence. Data were analyzed to estimate the prevalence of these conditions, to determine the demographic characteristics of the cases, and to describe the geographic distribution of these conditions within New Hampshire (Figure 1).

Mesothelioma was recorded on 13 death certificates, nine for males (average age at death: 65 years) and four for females (average age at death: 69 years). In eight cases, the mesotheliomas were pleural in origin; five occurred in the abdominal cavity. The average length of survival following diagnosis was 10 months for males and 4 months for females. For seven of the cases, occupational exposure to asbestos was likely to have occurred. The remaining cases occurred among persons whose primary occupations at time of death were recorded as housewives, a lawyer, and an insurance broker.

Silicosis was recorded as a cause of death on 22 death certificates (average age at death: 70 years); three of these were silicotuberculosis. All cases involved males employed in industries associated with a potential for exposure to silica dust.

Asbestosis was listed for nine deaths as the cause of death (average age at death: 74 years). The eight male workers were employed in trades with a potential for asbestos exposure; the female was identified as a housewife. Deaths associated with asbestos exposure (both mesothelioma and asbestosis) were clustered in geographic areas where the industrial use of asbestos is known to have occurred (Figure 1).

Anthracosis (i.e., coal workers' pneumoconiosis) was recorded as the cause of death for a coal miner and a crane operator. The death of one worker who was involved in the manufacture of woven belts was attributed to byssinosis (not usually classified as a pneumoconiosis), and three deaths were listed as "pneumoconiosis-unspecified."

An additional 218 death certificates indicated pulmonary fibrosis or interstitial fibrosis as a cause of death. Of these, 39 involved occupations with a high probability of past exposure to

Work-Related Illnesses — Continued

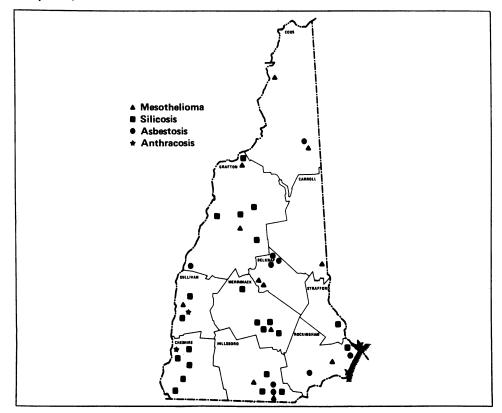
silica or asbestos (e.g., construction, shipbuilding, and manufacturing). Of the remainder, 15 cases involved work in the shoe or leather trades; 11 worked with wood or paper products; six were farmers; four worked with textiles; and four were painters; the remainder were unspecified.

Reported by E Schwartz, MD, State Epidemiologist, and staff, New Hampshire State Dept of Health and Human Svcs; National Institute for Occupational Safety and Health, CDC.

Editorial Note: NHDPHS elected to study deaths associated with mesothelioma and the pneumoconioses because these diseases are considered sentinel health events (occupational) (SHE[O]). A sentinel health event is defined as a preventable disease, disability, or untimely death, the occurrence of which serves as a warning signal that the quality of preventive activities and/or therapeutic medical care may need to be improved (2); a SHE(O) is an occupationally related sentinel disease. Mesothelioma and the pneumoconioses were specifically targeted for this study because: (1) the relationship is clearly established between these health conditions and exposure to dusts (e.g., asbestos, silica, coal) encountered in the workplace or general environment; and (2) recognition of a single case of these diseases justifies a careful search for other cases and raises important questions about their occurrence and the prevention of additional cases.

Mesothelioma is a tumor arising from the pleura or the peritoneum. A strong relationship has been established between the occurrence of mesothelioma and previous exposure to asbestos dust (3) among exposed workers, family contacts of asbestos-exposed workers, and

FIGURE 1. Death-certificate surveillance of selected work-related mortality — New Hampshire, 1963-1983



#### Work-Related Illnesses - Continued

persons incidentally exposed to airborne asbestos at work or from waste dumps, factories, construction sites, and automobile- or truck-brake linings. As with most chemically induced cancers, the latency period between the time of first exposure to asbestos and the clinical detection of mesothelioma is usually 20 years or more.

The pneumoconioses result from the inhalation and accumulation of dust in the lungs or from the reaction of the tissues to its presence (4). Although the inhalation of house dusts and most other dusts does not result in pneumoconiosis, the inhalation over long periods of time of silica, asbestos, and coal dust may result in pulmonary fibrosis. A pneumoconiosis may be suspected from the patient's occupational and medical history and is often diagnosed with the aid of chest radiographs and pulmonary-function tests.

For several reasons, this death certificate-based survey may have underestimated the magnitude of mortality associated with exposure to fibrogenic dusts. First, the long latency period usually associated with these diseases often hinders determination of their relationship to work. Accurate diagnosis requires that physicians consider the possible occupational or environmental origins of disease. A practical and systematic approach to such diagnoses has been devised (5). A key element is the occupational history, which should include details on various jobs held and information on chemical or physical exposures encountered during a patient's working lifetime (6). Second, because the clinical presentations of occupationally and nonoccupationally related diseases are often similar, it is not always clear whether certain diseases result from occupational exposure. Third, other potentially asbestos-related cancers (e.g., oropharyngeal, gastrointestinal, lung, or renal cancer) were not included in this survey. Fourth, because death certificates provide limited information on the decedent's occupational history, associations with specific occupations could not be determined for many of the cases. For example, approximately 10% of the death certificates listing pulmonary fibrosis or interstitial fibrosis as a cause of death also list occupation as "retired" or "at home." Fifth, based on the limited clinical and historic information on the death certificates, it is difficult to determine which cases of pulmonary fibrosis or interstitial fibrosis should be diagnosed as pneumoconioses. For the purposes of epidemiologic analysis, a pneumoconiosis was assumed if occupational information indicated exposure to asbestos or silica.

Investigation and follow-up of these sentinel health events may provide improved opportunity for prevention. Effective prevention of future pneumoconiosis and mesothelioma cases depends on: (1) recognition of work places where hazardous exposures are now occurring; (2) substitution of less hazardous substances for hazardous materials; (3) use of appropriate engineering controls, including local exhaust ventilation; (4) implementation of safe work practices; and (5) use of personal protective devices, such as respirators.

Effective surveillance and prevention depend on prompt reporting of pertinent diagnoses by those in the health-care community. Although in New Hampshire practicing health-care providers are required to report all cases of occupationally related disease to the state's Division of Public Health Services (7), relatively few cases are ever actually reported. It is essential, therefore, that health-care providers consider the potential occupational or environmental relatedness of diseases and report these diseases in conformance with reporting requirements.

#### References

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#### Work-Related Illnesses - Continued

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# Human T-Lymphotropic Virus Type III/ Lymphadenopathy-Associated Virus:

# **Agent Summary Statement**

#### INTRODUCTION

In March 1984, CDC and the National Institutes of Health (NIH), in consultation with scientists, physicians, and public health workers in academia, industry, and government, published a manual entitled *Biosafety in Microbiological and Biomedical Laboratories* ("biosafety manual")\* (1). The manual describes combinations of standard and special microbiologic practices, safety equipment, and facilities recommended for working with infectious agents in various laboratory settings. The recommendations are advisory and provide a voluntary code of safety practices.

A section of this manual is devoted to a number of specific "agent summary statements" consisting of brief descriptions of documented or anecdotal laboratory-associated infections, the nature of the laboratory hazards, and recommended precautions to be taken in handling and working with certain infectious agents. Contributors to the manual recognized that new agents would be discovered from time to time and recommended that a summary statement for each new agent be developed and published in the MMWR. The summary statement for human T-lymphotropic virus type III/lymphodenopathy-associated virus (HTLV-III/LAV)† follows. All laboratory directors are requested to put a copy of this summary in each of their copies of the biosafety manual and bring it to the attention of laboratory personnel. The recommendations in the summary statement were compiled from published scientific reports and are consistent with the published guidelines for health-care workers (2-4).

#### AGENT SUMMARY STATEMENT: HTLV-III/LAV

As of August 15, 1986, no cases of acquired immunodeficiency syndrome (AIDS) that meet the CDC case definition and can be attributed to an inadvertent laboratory exposure have been reported in laboratory workers (5). One laboratory worker (7) was included among the health-care workers who have had HTLV-III/LAV antibody detected in their serum after sustaining a needlestick injury (2,3,6-10), but the source of the infection could not be established. Persons who are infected with HTLV-III/LAV may be asymptomatic, may have AIDS-related complex, or may manifest symptoms of overt AIDS (11).

In 1985, two different reagent production laboratories reported that several laboratory workers may have been inadvertently exposed to an aerosol of concentrated HTLV-III/LAV; one worker was cut by a piece of glass from a broken carboy that contained HTLV-III/LAV-infected cells and culture fluid. None of the potentially exposed persons had shown evidence of seroconversion after 6 months in one incident and 12 months in the other as a result of these occupational exposures.

Other reports dealing with HTLV-III/LAV infection in health-care personnel, including

<sup>\*</sup>Available from Superintendent of Documents, U.S. Government Printing Office, Washington, D.C. 20402, Stock #01702300167-1, Price: \$4.00; and from National Technical Information Service, U.S. Department of Commerce, 5285 Port Royal Road, Springfield, Virginia 22161, Stock #PB84-206879, Price: \$6.00.

<sup>&</sup>lt;sup>†</sup>The Human Retrovirus Subcommittee of the International Committee on the Taxonomy of Viruses has proposed the name human immunodeficiency virus (HIV) for these viruses (Science 1986:232:697).

laboratory workers (3,4,6,8-10), indicate that the risk of bloodborne transmission from inadvertent exposure is considerably less for HTLV-III/LAV than for hepatitis B virus infection. These reports illustrate the need for complete evaluation by a physician and serologic testing of each laboratory worker definitely or possibly exposed to HTLV-III/LAV in a laboratory setting. It is recommended that the Public Health Service guidelines for health-care workers be followed in these instances (2.3).

#### Laboratory Hazards

HTLV-III/LAV has been isolated from blood, semen, saliva, tears, urine, cerebrospinal fluid, brain tissue, and cervical secretions and is likely to be present in other body fluids, secretions, and tissues of infected humans or experimentally infected nonhuman primates. Percutaneous or parenteral inoculation and direct contact of cuts, scratches, abrasions, or mucosal surfaces with suspensions of virus or specimens containing live virus are considered potential routes of infection. Possible transmission of infection via the parenteral route can occur through self-inoculation with needles, broken glass, or other sharp objects that contain HTLV-III/LAV. Spillage is a possible means of exposure and infection, especially spills accompanied by spraying or splashing of infected cell cultures, viral concentrates, and other infectious materials that may come into direct contact with abraded skin or mucous membranes of the eyes, nose, or mouth; however, there are no data documenting or suggesting that transmission of HTLV-III/LAV has occurred in this manner. Ingestion and inhalation have not been documented as modes of transmission of the virus.

#### **Recommended Precautions**

- 1. Biosafety Level (BSL) 2 standards and special practices, containment equipment, and facilities as described in the CDC-NIH biosafety manual are recommended for activities involving clinical specimens, body fluids, or tissues from humans or laboratory animals that may contain HTLV-III/LAV. These are the same practices recommended for all clinical specimens. Emphasis is placed on the following practices, which are included in the manual (1):
  - a. Use of syringes, needles and other sharp instruments should be avoided if possible. Used needles and cutting instruments should be discarded into a puncture-resistant container with a lid. Needles should not be resheathed, purposefully bent, broken, removed from disposable syringes, or otherwise manipulated by hand.
  - b. Gloves should be worn by all personnel engaged in activities that may involve skin contact with potentially infectious fluids, tissues, or cultures and by laboratory workers with dermatitis or other lesions on the hands who may have direct or indirect contact with potentially infectious materials. Handwashing with soap and water should be a routine practice immediately after direct contact with potentially infectious materials and on completion of work, even when gloves are worn.
  - c. Generation of aerosols, splashes, and spills of potentially infectious materials should be avoided in procedures involving body fluids or tissues, during necropsy of cadavers, and in similar procedures on animals experimentally infected with HTLV-III/LAV. Laboratory workers should use a biological safety cabinet when propagating the virus to further reduce the risk of exposure. Although the major precautions are listed here, the CDC-NIH biosafety manual contains additional related precautions (see pages 11-13 for BSL 2 and pages 14-17 [1] for BSL 3 when large volumes or concentrates of HTLV-III/LAV are involved). In all instances, the laboratory director is responsible for assessing the biosafety level to be used.
  - d. Human serum from any source that is used as a control or reagent in a test procedure should be handled at BSL 2 (see pages 11-13 [1]). Appended to this Agent Summary Statement is a statement (Addendum 1) issued by CDC on the use of all human control or reagent sera shipped to other laboratories. The Food and Drug Administration re-

quires that manufacturers of human serum reagents use a similarly worded statement.

- e. Animal BSL 2 practices, containment equipment, and facilities are recommended for activities involving nonhuman primates experimentally infected with HTLV-III/LAV. Laboratory coats, gowns, or uniforms should be worn by laboratory workers, as is customary for other BSL 2 or 3 practices, depending on the nature of the work, concentration of the virus, and volume of material being handled. Because many animals bite, and some throw feces, urine, or expectorate at humans, animal-care personnel must wear coats, protective gloves, coveralls or uniforms, and face shields as appropriate to protect the skin and mucous membranes of the eyes, nose, and mouth from potential exposure to these substances when working with animals likely to manifest such behavior.
- Activities such as growing research-laboratory-scale amounts of HTLV-III/LAV or related viruses or virus-producing cell lines, working with concentrated virus preparations, or conducting procedures that may produce droplets or aerosols should be performed in a BSL 2 facility with the additional practices and containment equipment recommended for BSL 3 (12).
- 3. Activities involving industrial-scale, large-volume, or high-concentration production and manipulation of HTLV-III/LAV are to be conducted with BSL 3 requirements (12).

(Continued on page 547)

TABLE I. Summary—cases specified notifiable diseases, United States

	3	4th Week Endi	ng	Cumulative, 34th Week Ending					
Disease	Aug. 23, 1986	Aug. 24, 1985	Median 1981-1985	Aug. 23, 1986	Aug. 24, 1985	Median 1981-1985			
Acquired Immunodeficiency Syndrome (AIDS)	176	136	N	8,113	4,955	N			
Aseptic meningitis	545	400	400	5,116	4,641	4,641			
Encephalitis: Primary (arthropod-borne	4								
& unspec.)	39	40	43	622	696	748			
Post-infectious	-	2	1	68	89	65			
Gonorrhea: Civilian	17,513	18,254	19,263	563,987	568,921	581,109			
Military	368	498	431	10,813	13,794	15,853			
Hepatitis: Type A	375	450	429	13,995	14,095	14,095			
Type B	524	527	480	16,728	16,485	15,407			
Non A, Non B	64	90	N	2,299	2,684	N			
Unspecified	57	110	129	2.994	3.715	4.662			
Legionellosis	15	13	N	412	464	N			
Leprosy	6	7	2	175	257	163			
Malaria	22	21	21	642	662	662			
Measles: Total*	64	32	17	5.186	2,332	2.215			
Indigenous	64	27	N	4.951	1,957	N.			
Imported	1 .	5	Ñ	235	375	N			
Meningococcal infections: Total	22	37	33	1,747	1,669	1,969			
Civilian	22	37	33	1.745	1.663	1,965			
Military	I	· ·	-	.,, 2	6	9			
Mumps	43	23	23	3,259	2.140	2.382			
Pertussis	97	158	53	1,926	1.672	1,344			
Rubella (German measles)	6	14	11	385	496	751			
Syphilis (Primary & Secondary): Civilian	806	526	603	17.106	17.353	19.654			
Military	1 000	2	5	110	118	240			
Toxic Shock syndrome	7	6	Ň	233	258	Ž-TŮ			
Tuberculosis	476	446	502	14.077	13.746	15.149			
Tularemia	10	5	7	89	115	156			
Typhoid fever	1 '5	6	13	180	223	257			
Typhus fever, tick-borne (RMSF)	39	20	25	525	451	743			
Rabies, animal	104	121	149	3,601	3,477	4.195			

TABLE II. Notifiable diseases of low frequency, United States

	Cum 1986		Cum 1986
Anthrax Botulism: Foodborne Infant (N.Y. City 1, Utah 3, Calif. 2) Other	6 36 1	Leptospirosis Plague Poliomyelitis, Paralytic Psittacosis (Mo. 1. Calif. 1)	23 4 - 69
Brucellosis (Fla. 1, Tex. 1, Colo. 1, Calif. 1) Cholera	51	Rabies, human Tetanus (Maine 1, Ohio 1, Mo. 1, Calif. 2)	-
Congenital rubella syndrome Congenital syphilis, ages < 1 year	107	Trichinosis (Ohio 1) Typhus fever, flea-borne (endemic, murine) (N.Y. City	43 21
Diphtheria	1.0%	1, Tex. 1)	34

<sup>\*</sup>There were no cases of internationally imported measles reported for this week.

TABLE III. Cases of specified notifiable diseases, United States, weeks ending August 23, 1986 and August 24, 1985 (34th Week)

		Aseptic	Encephalitis		Gon	н	epatitis (V	iral), by ty		Legionel-	l	
Reporting Area	AIDS	Menin- gitis	Primary	Post-in- fectious		vilian)	. А	В	NA,NB	Unspeci- fied	Legionel- losis  1986  15	Leprosy
	Cum 1986	1986	Cum 1986	Cum 1986	Cum. 1986	Cum. 1985	1986	1986	1986	1986		Cum 1986
UNITED STATES	8,113	545	622	68	563,987	568,921	375	524	64	57	15	175
NEW ENGLAND	355	25	17	3	14,152	15,040	10	41	-	5	-	6
Maine	13	2	-	-	590	727 376	:	2	-	-	•	-
N H Vt	8 3	3	2	2	353 165	203	1	2		-	- :	
V t Mass	187	6	4	-	5,757	5,749	3	28	-	5	-	6
RI	21	7	-	-	1,125	1,193	2	2	-	-	•	•
Conn	123	7	9	1	6,162	6,792	4	7	-	-	-	-
MID ATLANTIC Upstate N Y	3,112 296	31 4	68 26	6 4	95,430 11,291	82,762 11,054	9 3	27 13	1	13	9	12
N Y City	2,107	6	15	-	55,842	41,576	3	4	-	13	-	10
N J	492	21	10	-	12,446	12,467	3	10	. <del>.</del>			:
Pa	217	U	17	2	15,851	17,665	U	U	U	U	_	1
EN CENTRAL Ohio	489 100	109 38	177 56	10 2	73,788 19,317	76,449 19,586	21 9	35 14	8 3	4	1	4
ind	48	10	39	3	7,876	7,704	2	4	-	2	-	-
H	238	25	38	4	20,880	20,385	5	5	2	1		3
Mich	78	36	34	1	22,947	21,425	5	12	3	1	1	1
Nis	25	-	10	-	2,768	7,349	-	-	-	-	-	•
N N CENTRAL	160	26	29	8	24,294	26,399	8	21	3	-	1	2
∕linn owa	60 11	5 3	12 9	:	3,436 2,444	3,867 2,855	1	5 4	1	-	-	1
Mo	56	14	-	-	12,282	12,717	3	10		-		-
N Dak	2	-	-	-	214	181	-	-	-	-	-	-
S Dak	1	-	7		492	491	- :	:	- :	-	-	-
Vebr Cans	6 24	4	ī	1 7	1,824 3,602	2,255 4,033	3	2	1 1	-	1	1
ATLANTIC	1,120	69	82	23	147,114	146,239	32	98	13	4	1	2
Del	17	-	5		2,345	2,691	1	•	-	-	-	-
Λd	123	11	25	1	17,555	19,010	2	17	2	-	-	-
C	141	3		1	10,839	9,867	-	1	1		-	1
/a N Va	106 6	2	27 12	1	11,942 1,451	12,201 1,644	-	1	1	1		
I C	44	20	iī	1	22,717	22,046	2	23	2	2	-	
S C	24	-	-	-	12,807	14,025	3	9	-	-	-	•
ia Ia	170 489	8 25	2	1 18	24,802 42,656	29,631 35,124	3 21	10 33	1 6	ī	-	1
S CENTRAL	103	30	40	3	45,980	48,307	6	45	3	2	_	1
(y	21	15	19	1	5,055	5,446	2	9	-	-	-	-
Tenn .	53	4	3	1	17,753	18,471	1	16	1	2	-	-
Ala Viss	19 10	11	17 1	1	13,156 10,016	14,761 9,629	3	20	2	-	-	1
N S CENTRAL	488	206	81	6	67,952	71,757	100	98	16	21	2	16
Ark	21	-	-	2	6,204	7,021	6	3	2	1	-	i
.a	108	19	.3	-	12,078	14,154	18	32 9	3 1	3		
Okla Tex	27 332	16 171	16 62	4	7,559 42,111	7,725 42,857	15 61	54	10	17	-	15
MOUNTAIN	203	3	22	1	16,507	17,747	24	16	3	1	-	11
<b>M</b> ont	4	-	-	1	468	492 543	1 2	-	-	-	-	-
daho Nyo	2 4	1	2		536 362	420	-		-	-	-	-
Colo	96	1	3	-	4,193	5,230	1	6	1	1	-	3
N Mex	11	-	3	-	1,648	2,034	15	3	2	-	-	5
Ariz	50	-	8	-	5,364	5,205	2	2	-	-	-	1
Jtah Nev	12 24	ī	5 1	-	713 3,223	778 3,045	3	5	-	-	-	ż
PACIFIC	2,083	46	106	8	78,770	84,221	165	143	17	7	1	121
Wash	93	4	11	-	5,918	6,315	30	18	1	3	-	4
Oreg	44	- :		-	3,273	4,144	29	19	.4	-	-	
Calif	1,906	36 1	92	8	66,897	70,673	105 1	106	12	4	1	84
Alaska Hawaii	10 30	1 5	3	:	1,813 869	1,901 1,188	-	:	-	-	-	23
Guam		_	_		122	131	-		-		-	1
PR	76	4	4	-	1,539	2,262	3	9	-	2	-	7
					400	322	U	U	U			
VI. Pac Trust Terr	3	U	:	-	139 285	574	5		U	U 1	U	3 <b>3</b>

N Not notifiable

TABLE III. (Cont'd.) Cases of specified notifiable diseases, United States, weeks ending August 23, 1986 and August 24, 1985 (34th Week)

		August 23, 1986 and August 24, 1985 (34th Week)													
T	Malaria			sles (Rub			Menin- gococcal	Mu	mps	Γ	Pertussis	,	Γ	Rubella	
Reporting Area	Cum.	1986	Cum.	1986	Cum.	Total Cum.	Infections Cum.	-	Cum.	├—	Cum.	Cum.	1000	Cum	Cum
UNITED STATES	1986	L	1986		1986	1985	1986	1986	1986	1986	1986	1985	1986	1986	1985
NEW ENGLAND		64	4,951	-	235	2,332	1,747	43	3,259	97	1,926	1,672	6	385	496
Maine	33 1	:	75 10	-	8	123 1	124 23	:	53	2	105 2	83 5	-	9	12
N.H. Vt.	2	-	39	-	-	-	6	•	13	1	50	31	-	1	2
Mass	17	-	23	:	6	115	15 29	-	3 9	:	3 28	3 25	-	1	6
R.I. Conn.	5 7	-	2 1	-	2	7	16 35	:	9 19	-	4 18	12	-	2	4
MID ATLANTIC	90	47	1,606	_	21	197	273	-	134	4	133	104	_	31	205
Upstate N.Y. N.Y. City	30	2	64	-	19	82	93	-	53	4	86	61	-	23	17
N.J	25 18	45	615 905	•	2	60 27	57 29	:	5 36	-	.3	9	•	5	164
Pa.	17	Ū	22	Ū	-	28	94	Ū	40	Ū	11 33	31	Ū	3	11 13
E.N. CENTRAL Ohio	42	9	965	-	16	513	238	28	2,218	5	249	343	1	35	28
Ind.	12 2	6	17	:	10	54 57	94 19	1	100 31	5	108 22	32 70	-	1	1
ill.	14	-	631	-	3	289	66	27	1.644	-	28	36	-	24	12
Mich. Wis.	13 1	3	56 261	-	3	54 59	53 6	:	248 195	-	24 67	30 175	1	8	14
WN CENTRAL	22	1	322	_	17	11	84	_	82	14	163	94			
Minn. Iowa	5	-	45	-	4	6	17	-	1	'-	43	28	-	10	19
Mo.	1 10	1	133 25	-	1 6	2	11	-	21	-	13	5	-	1	1
N. Dak	-	-	25	-	1	2	28	-	15 3	-	12	24 9	-	1	7
S Dak. Nebr	4	-	-	-	-	-	4	-	1	-	14	ĭ	-		
Kans	2	:	94	-	5	1	9 15	-	41	14	1 76	4 23	-	7	7
S. ATLANTIC Del.	82	1	505	-	53	273	322	2	154	10	588	377	_	10	50
Md.	1 12	-	1 22	-	-	-	2	-	-	-	222	-	-		1
D.C.	1	-	- 22		9	88 8	44		15	•	136	226	-	:	6
Va. W. Va.	21	-	35	-	24	25	53	2	34	-	30	8	-	-	2
N.C.	4	-	2	-	1	33 9	3 56	-	38 14	3	23	2	-	-	9
S.C. Ga.	6	-	274	-		3	29	-	12	-	41 13	17 1		-	3
Fla.	7 26	1	79 90	:	14 3	8 99	49 82	-	14 27	7	102	76 47	:	10	29
E.S. CENTRAL	16	-	56	_	8	4	97	1	25	5	42	18		4	
Ky. Tenn.	4	-	_ :	-	6	2	24	-	6	-	5	3	-	4	2
Ala	1 7	-	54	-	1	1	35 27	1	16 2	3	15	6	-	-	
Miss.	4	-	2	-	:	1	11	-	1	2	22	6 3	-	-	
W.S. CENTRAL Ark.	63	-	585	-	34	423	153	4	151	30	165	249		55	32
La.	8		276 4	:	2	42	22 22	-	7 2	3 3	11 11	12 10	-	-	1
Okla. Fex	9		37	-	2	1	21	N	Ñ	-	89	120	-	-	1
	46	-	268	•	30	380	88	4	142	24	54	107	-	55	30
MOUNTAIN Mont	26	1	296	-	26 8	526 137	88 8	-	203 5	5 2	189	114	-	21	5
daho	1	-	1	-	-	135	3		6	-	10 33	7	-	2	1
Wyo. Colo.	-	-	-	-	-		. 2	-	-	-	1	-	-	-	
N. Mex.	8 4	-	2 32	-	5 7	13 5	13 8	N	11 N	1	52 17	32 11	-	1	2
Ariz.	8	-	252	-	6	236	19	- "-	167	-	46	27	-	2	1
Utah Nev.	2 3	1	8 1	:	-	-	9 26	-	10 4	2	27 3	30	-	13 3	i
PACIFIC	268	5	541		52	262	368	8	239	22	292	290	5	210	143
Wash.	21	-	158	-	25	42	54	-	7	1	82	52	-	14	11
Oreg. Calif.	15 231	1 3	4 358	:	4 22	3 197	26 275	N 8	N 209	-	10	29	-	. 1	1
Alaska	231		356	:	- 22	197	11	8	209	21	190 2	168 29	5	191	82
ławaii	1	1	21	-	1	20	2	-	17	-	8	12	-	4	48
Guam	1	-	4		1	11		-	4	-		-	-	3	2
P.R. 7.I.	4	.:	33	Ū	:	50 10	2	2	23	1	13	10	2	60	25
ac. Trust Terr.	-	U	-	-	:	10	1	U	13 7	U	-	:	U	2	:
Amer Samoa	-	-	2	-		-	:	-	4		-	-	-	ī	

<sup>\*</sup>For measles only, imported cases includes both out-of-state and international importations.

N Not notifiable U Unavailable

TABLE III. (Cont'd.) Cases of specified notifiable diseases, United States, weeks ending August 23, 1986 and August 24, 1985 (34th Week)

			Toxic-	na Augus			Γ	Typhus Fever	
Reporting Area	Syphilis (Primary &	(Civilian) Secondary)	shock Syndrome	Tuber	culosis	Tula- remia	Typhoid Fever	(Tick-borne) (RMSF)	Rabies, Animal
	Cum 1986	Cum 1985	1986	Cum 1986	Cum. 1985	Cum. 1986	Cum 1986	Cum 1986	Cum 1986
UNITED STATES	17,106	17,353	7	14,077	13,746	89	180	525 +	39 3,601
NEW ENGLAND Maine	308 15	352 9	-	451 32	467 35	1	11	9→	1 3
N.H	10	9	-	18	15	-	-	1	-
Vt Mass	7 165	5 175	-	13 228	4 277	1	9	2	-
RI	16	1/3	-	35	35		-	3	1
Conn	95	142	-	125	101	-	2	3	. 2
MID ATLANTIC	2,403	2,293	-	2,863	2,525	1	14	20 +	4 412
Upstate N Y N Y City	104 1,383	161 1,425		416 1,498	438 1,225	:	2 6	11 4	ł 54 -
NJ	433	447	-	497	349	1	5	1	15
Pa	483	260	U	452	513	-	1	3	343
E N CENTRAL	666 85	719 100	3 1	1,667 300	1,681 306	-	13 2	54 + 52 <b>2</b>	
Ind	78	65	÷	178	206	-	2	-	14
III Mich	351	362	-	733	738	-	2 5	1	26 20
Wis	117 35	148 44	2	378 78	325 106	-	2	-	21
W N CENTRAL	147	152	-	402	373	27	7	35+	7 584
Minn Iowa	27 6	31 17	-	101 33	79 43	i	1	1	82 130
Mo	80	74	-	199	177	22	5	18 6	63
N Dak	3	2	-	6	. 6	:	-	1 6 \	130
S Dak Nebr	11	5 7	-	16 7	18 13	2 1	-	6 \ 4	115 22
Kans	17	16	-	40	37	i	1	4	42
S ATLANTIC	5,384	5,119	1	2,710	2,797	8	24	246+	13 863
Del Md	35 291	25 309	:	27 209	27 256	2	1 6	28 2	432
DC	204	234	-	87	105	-	2		26
Va W Va	247 18	195 15	1	225 74	245 74	2	5 3	43	119 27
N C	334	437	-	372	368	i	3	87 €	5 7
s c	447	539	-	358	340	-	-	58 3	33
Ga Fla	985 2,823	888 2,477	-	408 950	454 928	3	4	21 1	132 81
ES CENTRAL	1,151	1,282		1,206	1,200	8	2		-5 <sub>236</sub>
Ky Tenn	53	42	-	287	269	3		16 5	
Ala	402 366	398 422		352 380	359 364	4 1	1 -	25 14	97 74
Miss	330	420	-	187	208	-	1	8	, 2
W S CENTRAL Ark	3,425 165	3,986 212	2	1,814 237	1,681 181	39 28	15	89 <del>1</del> 4 1	526 119
La	570	687	-	315	221	1	1		15
Okla Tex	87 2,603	116 2,971	2	166 1,096	176 1,103	6 4	1 13	74 <sup>4</sup>	45 347
MOUNTAIN	388	451	_	334	353	4	8	8	505
Mont	6	3	-	20	46	1	1	4	173
ldaho Wyo	9	4 6	-	14	15	-	-	i	2 221
Colo	1 96	112	-	28	5 43	-	1	3	15
N Mex	46	81	-	67	65	1	-	-	6
Arız Utah	158 12	219 5	-	160 28	147 10	1	3 2	-	79 3
Nev	60	21		17	22	i	ī	-	6
PACIFIC	3,234	2,999	1	2,630	2,669	1	86	1 4	1 382
Wash Oreg	99 75	80 60	-	123 92	150 85	-	3	-	5
Calif	3,033	2,810	1	2,250	2,241	:	79	ī )	369
Alaska Hawaii	2 25	2 47	-	37 128	68 125	1	1 3	- '	8
Guam		• • •	•			-	3		
P R	1 576	2 536	:	34 210	31 233	-	4		34
VI Pac Trust Terr	170	1 80	U	1 44	1 38		42	-	-

TABLE IV. Deaths in 121 U.S. cities.\* week ending August 23, 1986 (34th Week)

August 23, 1900 (34th Week)															
		All Caus	es, By A	ge (Yes	s)		P&I**		All Causes, By Age (Years)						
Reporting Area	All Ages	≥65	45-64	25-44	1-24	<1	Total	Reporting Area	All Ages	≥65	45-64	25-44	1-24	<1	P&I**
NEW ENGLAND	597	397	126	45	10	17	40	S. ATLANTIC	1,230	733	272	141	35	45	43
Boston, Mass.	161	91	43	15	5	6	18	Atlanta, Ga	122	72	27	17	5	1	1
Bridgeport, Conn	56	37	14	5	-	-	2	Baltimore, Md	182 89	118 53	39 17	22	1	2 6	3
Cambridge, Mass	21 22	15	4 5	2	-	-	1	Charlotte, N.C.	102	61	20	8 15	5 3	3	2 7
Fall River, Mass Hartford, Conn.	56	17 35	12	6	-	3	3	Jacksonville, Fla. Miami, Fla.	84	43	20	19	1	ĭ	Ź
Lowell, Mass	23	16	6	1	-	-	2	Norfolk, Va.	58	32	17	5	2	2	3
Lynn, Mass.	10	10	·		_			Richmond, Va.	79	39	18	11	3	8	7
New Bedford, Mass	s 22	15	3	3	-	1	- 1	Savannah, Ga.	36	24	6	3	2	1	1
New Haven, Conn.		28	9	4	1	-	2	St. Petersburg, Fla.	104	86	13	2	1	2	5
Providence, R.I.	53	34	9	5	-	5	5	Tampa, Fla	67 283	32 162	18 69	6 32	3 5	15	7
Somerville, Mass.	8 27	.8	-	-	:		1	Washington, D.C.	263	111	8	1	4	13	′.
Springfield, Mass. Waterbury, Conn.	33	21 25	3 3	3	1 2	1	2	Wilmington, Del		• • •	·		-		
Worcester Mass	63	45	15	1	í	1	4	E.S. CENTRAL	720	461	144	75	28	12	25
	•••	70		•	•		- 1	Birmingham, Ala	116	71	23	15	4	3	2
		1,658			73	78	102	Chattanooga, Tenn.	47	33	. 7	7	-	-	5
Albany, N.Y.	55	32	14	6	2	1	2	Knoxville, Tenn	73	45	17	7	3	1	5
Allentown, Pa Buffalo, N.Y.	41 112	31 80	10 22	6	4	-	5	Louisville, Ky	85 159	61 97	12 39	7 13	2 10	3	6 3
Camden, N.J.	49	29	13	2	3	2	°.	Memphis, Tenn. Mobile, Ala	78	57	13	6	1	ī	3
Elizabeth, N.J	29	19	7	3	3	-	- 1	Montgomery, Ala.	59	35	12	9	3		2
Erie, Pa.†	36	20	12	-	1	3	1	Nashville, Tenn.	103	62	21	11	5	4	2
Jersey City, N.J.	41	19	10	7	2	3	-								
	1,339	817			33	39	51	W.S. CENTRAL	1,301	741	293	145	68	54	52
Newark, N.J.	71	39	11	7	5	6	3	Austin, Tex.	53	34	8	6	2	3	6
Paterson, N.J.	37	25	8	4	.:		2	Baton Rouge, La	37	27	6	2	-	2	3
Philadelphia, Pa Pittsburgh, Pa †	393 62	244 39	94 16		11	15	18	Corpus Christi, Tex. Dallas, Tex.	20 192	13 96	3 51	2 21	.2		1
Reading, Pa.	24	22	10	3	1	2	3 2	El Paso, Tex	53	27	16	3	12	12 3	3
Rochester, N.Y.	105	78	19	3	3	2	7	Fort Worth, Tex.	88	50	14	11	6	7	4
Schenectady, N.Y.	32	27	2	ĭ	ĭ	ī	il	Houston, Tex	287	136	84	42	15	10	7
Scranton, Pa.†	22	17	3	-	2	-	- 1	Little Rock, Ark	100	54	25	12	4	5	5
Syracuse, N.Y.	87	63	17	2	2	3	2	New Orleans, La.	119	80	22	13	3	1	1
Trenton, N.J.	28	17	7	3	:	1	1	San Antonio, Tex.	185	113	38	17	12	5	10
Utica, N.Y. Yonkers, N.Y.	19 29	17 23	1 6	-	1	-	1 3	Shreveport, La. Tulsa, Okla.	62 105	39 72	10 16	5 11	5 3	3	3 5
				-	-	-	۱	TOISO, ONIO	.00	′-		• • •	3	3	5
	2,294	1,477			65	74	89	MOUNTAIN	605	372	127	60	21	22	25
Akron, Ohio	43	27	9	6	1	-	-	Albuquerque, N.Me:	k 90	51	19	11	3	4	2
Canton, Ohio	26	17	7	1	• •	1	.5	Colo Springs, Colo	38	26	.7	5	-	-	7
Chicago, III.§ Cincinnati, Ohio	564 195	362 141	125 36	45 8	10 7	22 3	16 18	Denver, Colo Las Vegas, Nev	109 81	66 47	26 22	11 9	3 2	3	-
Cleveland, Ohio	147	96	38	6	ź	5	2	Ogden, Utah	19	14	1	2	1	i	3
Columbus, Ohio	127	80	30	10	6	ĭ	ī	Phoenix, Ariz	125	71	30	9	8	ż	4
Dayton, Ohio	96	65	20	8	2	1	3	Pueblo, Colo	17	15	1	ĭ	-		
Detroit, Mich.	262	146	55	46	10	5	9	Salt Lake City, Utah	39	20	6	7	2	4	1
Evansville, Ind	41	27	10	1	1	2	2	Tucson, Ariz	87	62	15	5	2	3	6
Fort Wayne, Ind.	69	49	11	5	3	1	-	DACIFIC	4 705						
Gary, Ind. Grand Rapids, Micl	10 h 55	5 39	2 10	2 4	ī	1	9	PACIFIC Berkeley, Calif.	1,735	1,134	320	169	66	39	80
Indianapolis, Ind	172	97	47	12	9	1 7	1	Fresno, Calif.	24 64	19 <b>42</b>	5 11	7	ī	3	4
Madison, Wis	55	35	8	6	4	ź	6	Glendale, Calif	21	19	' i	í	'.	3	1
Milwaukee, Wis.	129	85	23	14	-	7	ž	Honolulu, Hawaii	77	50	14	6	6	1	4
Peoria, III.	53	35	8	4	3	3	4	Long Beach, Calif.	95	55	25	9	ĭ		6
Rockford, III.	32	24	3	4	-	1	1	Los Angeles, Calif.	455	294	87	46	17	5 7	21
South Bend, Ind.	51	36	8	2	3	2	3	Oakland, Calif.	74	46	11	11	3	3	4
Toledo, Ohio	117	73	27	8	3	6	2	Pasadena, Calif.	31	22	6	. 1	:	2	1
Youngstown, Ohio	50	38	5	4	-	3	-	Portland, Oreg. Sacramento, Calif.	120	78	24	13	4	1	:
W.N. CENTRAL	671	449	147	36	24	15	24	San Diego, Calif.	121 130	80 75	25 24	7	4	5	5
Des Moines, Iowa	67	48	10	4	4	1	1	San Francisco, Cali		101	21	12 32	13 1	3	9 5
Duluth, Minn.	29	18	7	2	1	i	i	San Jose, Calif.	149	94	35	10	7	3	9
Kansas City, Kans.		31	13	3	1	-	2	Seattle, Wash.	131	96	19		6	3	1
Kansas City, Mo.	134	92	34	4	2	2	6	Spokane, Wash	48	36	. 5		3	-	2
Lincoln, Nebr.	24	19	. 1	2	1	1	1	Tacoma, Wash.	39	27	7		-	2	4
Minneapolis, Minn		25	14	6	2	1	2	LTOTAL						_	
Omaha, Nebr. St. Louis, Mo.	69 146	44	16	2	5	2	1	TOTAL	11,764	<sup>††</sup> 7,422	2,460	1,117	390	356	480
St. Louis, Mo. St. Paul, Minn.	146 55	102 37	33 9	5	2	4	3	1							
Wichita, Kans.	55 51	37	10	5 3	4	2	2 5	1							
**************************************	51	33	10	3	-	•	9	I							

<sup>•</sup> Mortality data in this table are voluntarily reported from 121 cities in the United States, most of which have populations of 100,000 or more. A death is reported by the place of its occurrence and by the week that the death certificate was filed. Fetal deaths are not included "Pneumonia and influenza.

<sup>†</sup> Because of changes in reporting methods in these 3 Pennsylvania cities, these numbers are partial counts for the current week Complete counts will be available in 4 to 6 weeks.

<sup>††</sup>Total includes unknown ages. § Data not available. Figures are estimates based on average of past 4 weeks.

- 4. All laboratory glassware, equipment, disposable materials, and wastes suspected or known to contain HTLV-III/LAV must be decontaminated, preferably in an autoclave, before washing, discarding, etc. Incineration of solid wastes may be used as an alternate method of disposal.
- 5. There is no evidence that laboratory clothing soiled with materials known or suspected to contain HTLV-III/LAV poses a transmission hazard, and the handling of such clothing is covered under BSL 2 practices. However, to be consistent with BSL 3 recommendations (1), when laboratory clothing becomes contaminated with HTLV-III/LAV preparations, it should be decontaminated before being laundered or discarded.
- 6. Work surfaces should be decontaminated at the end of each day on completion of procedures or when overtly contaminated. Many commonly used chemical disinfectants with such active ingredients as sodium hypochlorite, formaldehyde, glutaraldehyde, or phenols (4,13-15) can be used to decontaminate laboratory work surfaces; they can also be used to decontaminate some laboratory instruments, specific areas of contaminated laboratory clothing, and spills of infectious materials. Prompt decontamination of spills and other overt contamination should be standard practice.
- 7. The prudent and recommended approach to handling human serum known or suspected to contain HTLV-III/LAV is to use the same precautions that should be used routinely to prevent transmission of bloodborne infections, including hepatitis B (16). Available data on the effectiveness of heat to destroy HTLV-III/LAV suspected or known to be present in human serum are at variance because of variations in volume of serum, concentration of the virus, temperature, and duration of exposure to heat (14,15,17). Similarly, results of chemical analyses or antibody assays may vary when sera are heated before testing according to the analysis or assay being performed (18-20). However, there is agreement that testing heated serum for HTLV-III/LAV antibody by enzyme immunoassays often yields false-positive results (21-23).
- 8. No HTLV-III/LAV vaccine has been developed, and no drugs have been shown to be safe and effective for therapy. As part of an ongoing medical surveillance program for employees, all laboratory workers before being assigned to activities with a high potential for exposure should have a serum sample obtained and stored at -40 C (-40 F) for possible future testing. Subsequent serum samples should be obtained and stored in accordance with laboratory policy or following an inadvertent laboratory exposure involving materials described above. When indicated, these serum specimens should be tested by a qualified laboratory using currently recommended procedures for HTLV-III/LAV antibody. Furthermore, the physician requesting serologic testing of these serum specimens must first obtain informed consent from the laboratory worker and describe the confidentiality safeguards available to protect test results. The laboratory workers whose serum specimens are to be tested should understand how the test results are to be used, the implications of a positive or negative test result, and the limits, if any, of the confidentiality safeguards. An employee whose serum HTLV-III/LAV antibody test is reactive and whose subsequent tests and evaluation confirm the presence of HTLV-III/LAV infection should be counseled to follow the Public Health Service recommendations for preventing transmission (24,25).
- 9. In addition to HTLV-III/LAV, other primary, as well as opportunistic, pathogenic agents may be present in the body fluids and tissues of persons who are antibody positive or have AIDS-related complex or AIDS. Laboratory workers should follow accepted biosafety practices to ensure maximum protection against inadvertent laboratory infection with agents other than HTLV-III/LAV that may also be present in clinical specimens.

Reported by Div of Safety, National Institute of Allergy and Infectious Diseases, National Cancer Institute, National Institutes of Health; AIDS Program, Hospital Infections Program, Center for Infectious Diseases, Laboratory Program Office, Office of Biosafety, Office of the Director, CDC.

#### **ADDENDUM**

CDC cautionary notice for all human serum samples used as controls or reagents:

WARNING: Because no test method can offer complete assurance that laboratory specimens do not contain HTLV-III/LAV, hepatitis B virus, or other infectious agents, this specimen(s) should be handled at the BSL 2 as recommended for any potentially infectious human serum or blood specimen in the CDC-NIH manual, *Biosafety in Microbiological and Biomedical Laboratories*, 1984, pages 11-3.

One or more of the following statements should be included with the above warning statement:

- This specimen is negative for hepatitis B surface antigen (HBsAg).
- This specimen is negative for antibody to HTLV-III/LAV.
- This specimen is positive for hepatitis B surface antigen (HBsAg).
- This specimen is positive for antibody to HTLV-III/LAV.
- This specimen has NOT been tested for hepatitis B surface antigen (HBsAg).
- This specimen has NOT been tested for antibody to HTLV-III/LAV.
- This specimen has been heated at 56 C (133 F) for 30 minutes (which will not inactivate HBsAg but will inactivate HTLV-III/LAV).

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# Thrombotic Thrombocytopenic Purpura Associated with *Escherichia coli* O157:H7 — Washington

Since Escherichia coli was first associated with bloody diarrhea in 1982 (1), several well-documented cases of hemolytic-uremic syndrome (HUS) related to E. coli O157:H7 have been described among children and adults (2,3). Patients with thrombotic thrombocytopenic purpura (TTP) have clinical and pathologic features similar to patients with HUS. In contrast to HUS, however, gastrointestinal infections have not been strongly implicated in the pathogenesis of TTP (4,5). A patient who recently died in Seattle with a clinical and pathologic diagnosis of TTP had bloody diarrhea associated with E. coli O157:H7 infection for 1 week before the onset of her other symptoms. This patient's clinical course suggested that E. coli O157:H7 infection may have been related to the development of TTP.

In February 1986, a 53-year-old, previously healthy woman was admitted to a Seattle hospital with severe neurologic abnormalities, microangiopathic hemolytic anemia, throm-bocytopenia, and renal failure. One week before admission, she had developed bloody diarrhea. There was no past history of bowel disease or travel. The patient had not eaten hamburger or shellfish within the preceding 2 weeks, and in the several days before onset of diarrhea, she had eaten the same foods as her husband who did not become ill. Three days after onset of diarrhea, a hematocrit was normal, and a stool examination for ova or parasites was negative. Cephalexin and metronidazole were prescribed. Her diarrhea continued, and hematuria developed. The day before admission, she became extremely lethargic. Following a generalized seizure while being evaluated in an emergency room, she was admitted.

On admission, the patient was intubated and comatose. Rectal temperature was 35.6 C (96.1 F); blood pressure, 180/100 mm Hg supine; heart rate, 66 beats per minute; and respiratory rate, 10/minute on the ventilator. Pupils were pinpoint and minimally reactive. Cranial nerve functions were intact, and the patient withdrew her extremities to noxious stimuli. The abdomen was soft without guarding and bowel sounds were hypoactive. Stool was guaiac positive. Laboratory data included a hematocrit of 41%, a leukocyte count of 29,200/mm³ with a left shift, and a platelet count of 38,000/mm³. A blood smear showed schistocytes, helmet cells, burr cells, and target cells consistent with a microangiopathic hemolytic anemia. The serum urea nitrogen was 48 mg/dl, and creatinine was 3.3 mg/dl. PT and PTT were normal, and the thrombin time was 22 seconds. Fibrinogen was 454 mg/dl, and fibrin degradation products were less than 40  $\mu$ g/ml but greater than 10  $\mu$ g/ml. A culture from a rectal swab obtained the day of admission yielded *E. coli* 0157:H7.

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TTP was diagnosed; plasmapheresis was begun after treatment with fresh frozen plasma, and she received corticosteroids, dipyridamole, and aspirin. Her neurologic status deteriorated rapidly during the first 24 hours after admission, and a computerized tomographic scan of her head showed diffuse cerebral edema. Despite aggressive therapy for increased intracranial pressure, she lost all cranial nerve and motor function and died 48 hours after admission. An autopsy revealed diffuse fibrin microthrombi in multiple organs, including kidneys, pituitary, and myocardium. Transtentorial herniation appeared to be the cause of death. Cultures of food samples from the patient's refrigerator yielded no pathogens.

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Editorial Note: The syndromes of TTP and HUS share many features, including thrombocytopenia, microangiopathic hemolytic anemia, and renal disease (4,6). Neurologic manifestations and fever complete the pentad of symptoms characteristic of TTP; these symptoms can also be present in HUS. Diarrhea, frequently bloody, is usually present a few days before the onset of HUS. Although diarrhea is not often described before onset of TTP, abdominal pain has been noted in 13%-14% of patients (4,7), and major gastrointestinal hemorrhage can occur (7).

Verotoxin-producing *E. coli*, including *E. coli* O157:H7, has been associated with HUS (2,3,8). It has been postulated that one or more verotoxins may be involved in HUS, but the mechanism remains unknown. A rise in verotoxin-neutralizing antibody has been detected in paired sera from patients with HUS (2). Drugs, toxins, infectious agents, immunologic disease, and pregnancy have all been proposed as etiologic agents for TTP (4). The isolation of *E. coli* O157:H7 from stool of the patient described here with bloody diarrhea and TTP suggest infection with this agent may represent another feature common to both TTP and HUS. *E. coli* O157:H7 can cause nonbloody diarrhea and asymptomatic infection (8,9). It is not known whether *E. coli* O157:H7 may be associated with HUS or TTP without antecedent bloody diarrhea.

Recovery of *E. coli* O157:H7 is optimal when specimens are obtained within 6 days after onset of diarrhea (10). Laboratories can screen for *E. coli* O157:H7 using media containing sorbitol, since *E. coli* O157:H7 does not ferment sorbitol rapidly, whereas 93% of other *E. coli* do (11). Isolates that do not ferment sorbitol within 24 hours can be sent to an appropriate reference laboratory for serotyping. However, since *E. coli* other than serotype O157:H7 have also been associated with HUS (2), and these *E. coli* may ferment sorbitol, a thorough investigation requires screening of stool specimens by a reference laboratory. An aliquot of stool can be frozen at -70 C (-94 F) as soon as the diagnosis of HUS or TTP is considered, pending the investigation of other causes. CDC's Enteric Diseases Branch is interested in obtaining frozen stool specimens from patients with HUS or TTP to analyze for verotoxin-producing *E. coli*; arrangements can be made through state epidemiologists or laboratory directors.

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# Aseptic Meningitis Among Kidney Transplant Recipients Receiving a Newly Marketed Murine Monoclonal Antibody Preparation

During July 1986, four patients being treated for kidney allograft rejection in two hospitals developed headache, fever, and photophobia (three patients) or fever alone (one patient) during administration of a recently marketed murine monoclonal antibody preparation, Orthoclone OKT3\* (Ortho Pharmaceutical Corporation, Raritan, New Jersey). All patients had onset of symptoms within 72 hours of beginning their course of treatment with the product. OKT3 had been administered daily as 5 mg OKT3 mixed with 50 ml normal saline and infused intravenously over a 1-hour period. Lumbar puncture results on all four patients suggested aseptic meningitis; they revealed cerebrospinal fluid (CSF) pleocytosis, elevated CSF protein, and normal CSF glucose. Initial CSF white blood cell counts ranged from 18 to 445/mm³, with 40%-89% polymorphonuclear neutrophils. All bacterial cultures of CSF and blood were sterile. Viral cultures of throat and stool were done in two patients and CSF in one patient; all have remained negative. Symptoms in all four patients resolved within 5 days with no evidence of neurologic sequelae. Two of the patients received neither antibiotic nor antiviral therapy.

Four other patients who were treated for kidney allograft rejection during July 1986 at the same hospitals but did not receive OKT3 had no signs or symptoms suggesting possible aseptic meningitis. All eight patients received other drug and therapeutic interventions commonly given to treat kidney allograft rejection.

Investigations of the reactions are currently under way by hospital, state, Food and Drug Administration (FDA), and company officials.

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Editorial Note: OKT3 has been reported effective in the treatment of acute kidney allograft rejection (1). Commonly reported adverse reactions have included fever, chills, dyspnea, chest pain and tightness, wheezing, nausea and vomiting, and tremor (1). In one study, fever was reported in up to 73% of recipients. Reactions were noted 45-60 minutes after the first injection; second injections were associated with fewer reactions, and symptoms were uncommon in subsequent injections (1). The incidence of aseptic meningitis during treatment with OKT3 is not known. Sixteen (2%) of 977 patients who received OKT3 during premarket trials had a lumbar puncture. Complete information was available for only seven of the 16 pa-

<sup>\*</sup>Use of trade names is for identification only and does not constitute endorsement by the Public Health Service, U.S. Department of Health and Human Services.

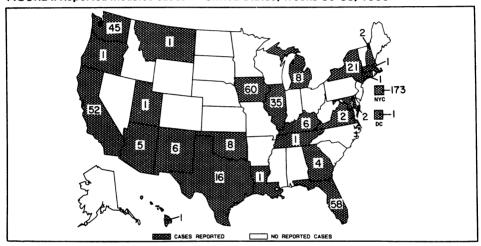
## Aseptic Meningitis — Continued

tients. In six of the seven patients, CSF cell count, CSF protein, and CSF glucose were consistent with aseptic meningitis; all bacterial, fungal, and viral cultures were negative. The extent and nature of meningeal pathology is unknown. Micrologic evaluation is incomplete for all patients but suggests an immunologically mediated reaction as one possible etiology. No clinical sequelae directly attributable to the reactions have been identified.

To assess the occurrence of aseptic meningitis and to establish incidence rates in patients receiving OKT3, the FDA and Ortho Pharmaceuticals have instituted a postmarket clinical evaluation of the product. The company has provided additional information about this evaluation to potential users of their product. Other physicians interested in this information should contact Ortho Pharmaceuticals, telephone (800) 433-1015; in New Jersey, (201) 218-6410. Reference

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FIGURE I. Reported measles cases — United States, weeks 30-33, 1986



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